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EXPLORING A NEW HYPOTHESIS FOR THE RAPID RISE OF COLON CANCER IN JAPAN AND SINGAPORE: REDUCTION OF IMMUNOSURVEILLANCE BY THE LOSS OF SYMBIOTIC APICOMPLEXA PROTOZOA. *D A Juckett, C Fylsworth, J M Quensen, and B Rosenberg. (Barros Research Institute, Holt, MI 48842)

The rapid rise in colon cancer within modernizing countries has not been adequately explained although several risk factors related to diet have been proposed and extensively explored in this and other settings. In a recent paper (Rosenberg et al., Int. J. Canc., DOI: 10.1002/ijc.20801), we showed that the presence of the endemic *Eimeria* spp. protozoan in intestinal tissues associated with regions of low tumorigenesis in the large and small ovine intestine and that an *Eimeria* surface protein is a potent activator of dendritic cells and a useful immunostimulant. We suggest that it may also enhance immunosurveillance by elevating the intestinal alert status. Since colon cancer is rare among vertebrates harboring *Eimeria*, the unusually high incidence in select human populations may be due to the elimination of this and related protozoa by modern sanitation. To determine if this hypothesis warrants investigation, a preliminary exploration of the ecological data for Japan and Singapore populations was undertaken. We show that colon cancer in these countries began to rise before changes in dietary patterns, especially in older populations less likely to alter their diets. This violates the tenet of causality. In contrast, the cessation of agricultural use of urban nightsoil precedes the rise in colon cancer. This change in fertilizer practice could break the protozoa's lifecycle and lead to the loss of this potential symbiont. We present both incidence and mortality time trends and comparisons to sanitation and lifestyle indicators. We conclude that this hypothesis is worth further consideration.

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DIETARY PHYTOESTROGEN INTAKE IS ASSOCIATED WITH REDUCED COLORECTAL CANCER RISK. *M Cotterchio, B Boucher, M Manno, S Gallinger (Cancer Care Ontario, Toronto, ONT Canada)

Phytoestrogens may reduce the risk of certain hormone-mediated cancers. Although this relationship has been most extensively studied with breast and prostate cancers, a possible protective role with colorectal cancer has also been suggested; though a paucity of studies have evaluated this. Phytoestrogens are plant compounds with non-steroidal estrogen-like activities. Important sources include isoflavones which are mostly found in legumes (eg., soy) and lignans which are widely distributed in grains, nuts, fruits, and vegetables. Isoflavones have dominated phytoestrogen research since they are major dietary components in Asia where hormone-dependent cancers are less prevalent. However, lignans are the most abundant phytoestrogen in North American diets therefore it is important to measure total phytoestrogen intake, and only recently has this been attempted by modifying food frequency questionnaires and analytic databases. We conducted a colorectal cancer case-control study. Cases were diagnosed with colorectal cancer, between 1997 and 2000, aged 20-74 years, identified through the population-based Ontario Cancer Registry and recruited by the Ontario Familial Colorectal Cancer Registry. Controls were a sex- and age-matched random sample of the population of Ontario. Approximately 1120 cases and 1900 controls completed epidemiologic and diet food frequency questionnaires. Multivariate logistic regression analysis was used to obtain adjusted odds ratios (OR) and estimates. We found that lignan intake was associated with a significant reduction in colorectal cancer risk (OR (T1 vs T3) = 0.73; 95% CI: 0.56, 0.94). Isoflavone intake was also significantly associated with colorectal cancer risk (OR (T1 vs T3) = 0.77; 95% CI: 0.58, 0.86). This finding that phy-

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HORMONE REPLACEMENT THERAPY AND COLORECTAL ADENOMA: DATA FROM THE PROSTATE, LUNG, COLORECTAL AND OVARIAN (PLCO) CANCER SCREENING TRIAL. *M P Purdue, P J Mink, P Hartge, W-Y Huang, J Weissfeld, S Buys, R B Hayes (Division of Cancer Epidemiology and Genetics, National Cancer Institute, Rockville, MD 20892)

PURPOSE: Findings from some epidemiologic studies of colorectal cancer and adenoma in women suggest that the protective effect of hormone replacement therapy (HRT) may differ across categories of post-menopausal age and body mass index (BMI). We conducted an analysis of women participating in the Prostate, Lung, Colorectal, and Ovarian (PLCO) Cancer Screening Trial to investigate the relationship between HRT use and prevalent adenoma, with assessment of age and BMI subgroup-specific risks. **METHODS:** We identified 1,468 women with at least one left-sided adenoma and 19,203 women without adenoma or colorectal cancer. Information about HRT and reproductive factors was obtained from a self-administered questionnaire. **RESULTS:** Compared to never use of HRT, current use was associated with a decreased prevalence of left-sided adenoma (odds ratio (OR) 0.85; 95% confidence interval 0.75-0.97). We found no evidence of dose-response with increasing duration of use for current or former users. The association with current HRT use was stronger among women aged 65+ (OR 0.94, 0.69 among women aged <65 and 65+ respectively; $P_{\text{interaction}} = 0.03$) and among women with a BMI < 30 (OR 0.83, 0.95 among women with BMI < 30, 30+ respectively; $P_{\text{interaction}} = 0.40$). Reproductive factors were not significantly associated with adenoma prevalence. **CONCLUSION:** We found a reduced prevalence of adenoma with current HRT use, particularly among older women.

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ASSOCIATION BETWEEN COLONIC SCREENING, SUBJECT CHARACTERISTICS AND STAGE OF COLORECTAL CANCER. *L Fazio, M Cotterchio, M Manno, J McLaughlin, S Gallinger (Cancer Care Ontario, Toronto, Ontario M5G 2L7)

Colorectal cancer survival is highly dependent on stage at diagnosis. This study evaluated the association between subject factors and stage of colorectal cancer at diagnosis. Population-based cases recruited by the Ontario Familial Colon Cancer Registry (OFCCR) diagnosed between 1997 and 1999 were staged according to the tumor-nodal-metastasis (TNM) staging system and classified as early (TNM I/II) or late (TNM III/IV) stage. Multivariate logistic regression was used to determine associations between subject factors and late (versus early) stage colorectal cancer. Among 768 cases, 60% were early and 40% were late stage. The most significant predictors of late stage in the multivariate model were: age at diagnosis compared to <45 years of age: (45-59: OR = 0.36 95% CI 0.18-0.74 and (60+: 0.30 95% CI 0.15-0.60), rural residence (OR = 1.5; 95% CI 1.0 - 2.17), non-white race (OR = 3.3; 95% CI 1.2 - 9.3), first degree relative with colorectal cancer (OR = 0.7; 95% CI 0.5 - 0.9), and screening endoscopy (OR = 0.5; 95% CI 0.2 - 1.0). Increased vegetable consumption was associated with a decreased risk of late stage cancer, and was of borderline significance. In the adjusted analysis age > 45 years, a first degree relative with colorectal cancer and screening endoscopy were associated with early (versus late) stage and rural residence and being non-white was associated with late (versus early) stage. Colorectal cancer screening and awareness programs are likely to present with late stage cancer.